## Avian Influenza A (H7N9) Key Points June 7, 2013

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#### **Situation Summary**

- New reports of human infections with avian influenza A (H7N9) have decreased since the month of April, when multiple cases were being reported each day.
- As of May 30, 2013, 132 human cases of H7N9 infection and 37 deaths <u>have been</u> reported to World Health Organization.
- The decline in newly reported cases may be a result of control measures implemented in China – including closure of live bird markets – or it may be a result of a seasonal pattern previously seen with other avian influenza viruses. (For more information, see "H7N9 Situation and CDC Response Update" at <a href="http://www.cdc.gov/flu/spotlights/h7n9-cdc-response.htm">http://www.cdc.gov/flu/spotlights/h7n9-cdc-response.htm</a>.)
- The first human cases of H7N9 virus infection were internationally reported by WHO on April 1, 2013.
- No cases of H7N9 have been detected in the United States or anywhere outside of China at this time.
- Most of the reported cases of human infection with H7N9 have had very serious illness.
- Mild illness or asymptomatic illness has been reported rarely among some people with H7N9 virus infection.
- The H7N9 situation is of international public health concern because of the potential for this virus to change and trigger a pandemic, which could be severe based on the epidemiological information currently available.
- An ongoing investigation suggests that most people have been infected with the
  virus after having contact with infected poultry or contaminated environments. (See
  "Avian Influenza Background" for information on how this type of transmission might
  occur.)
- Follow-up investigations by Chinese health officials of more than 2,000 close contacts of people infected with H7N9 indicate that there is no sustained (ongoing) spread of this virus from person to person at this time; however a few small clusters of human

infection have occurred where the possibility of limited human-to-human spread cannot be excluded.

- As noted in the Health Update CDC distributed via the Health Alert Network on June 7, 2013, preliminary results from influenza-like illness surveillance suggest that H7N9 has not caused widespread human illness in China.
- Based on previous experience with other avian influenza viruses, it would not be surprising to see some limited human-to-human spread of this H7N9 virus.
- Most important, however, is the fact that there is no evidence of efficient and sustained (ongoing) spread in the community, which is what is needed to trigger a pandemic.
- Although there is no evidence of sustained person-to-person transmission, CDC is conducting routine pandemic preparedness activities.
- A candidate vaccine virus has been developed in case it should be needed. CDC is working now to evaluate and prepare additional potential H7N9 candidate vaccine viruses. (See section H7N9 Candidate Vaccine Viruses below).
- CDC also is examining available H7N9 viruses for susceptibility to the licensed antiviral drugs, oseltamivir and zanamivir (Read more about H7N9 and antiviral drugs in the section below).
- On June 7, 2013, CDC issued a Health Update via the Health Alert Network (HAN) that provides a summary of the H7N9 situation and includes new recommendations on who should be tested for H7N9 in the United States. The Health Update is available at <a href="http://emergency.cdc.gov/HAN/han00347.asp">http://emergency.cdc.gov/HAN/han00347.asp</a>
- CDC also has posted Interim Guidance for Specimen Collection, Processing, and Testing for Patients Who May Be Infected with Avian Influenza A (H7N9) Virus. The guidance is available at <a href="http://www.cdc.gov/flu/avianflu/h7n9/specimen-collection.htm">http://www.cdc.gov/flu/avianflu/h7n9/specimen-collection.htm</a>
- The H7N9 case definitions have been updated. The Interim Guidance on Case Definitions to be Used for Avian Influenza A (H7N9) Case Investigations in the United States is available at <a href="http://www.cdc.gov/flu/avianflu/h7n9/case-definitions.htm">http://www.cdc.gov/flu/avianflu/h7n9/case-definitions.htm</a>

#### What May Happen

- It would NOT be surprising to see **limited** human-to-human transmission during the current H7N9 situation in China.
- It's also possible that H7N9 cases may at some point be detected in the United States (for example, in a traveler returning from China).
- This would not signal an increase in the potential risk to the public's health unless the transmission pattern of the virus was to change.
- If a person in the United States has H7N9 flu, they will be isolated (separated from other people who are well) and cared for.
- While there is no vaccine against this virus at this time, influenza antiviral drugs can be used for treatment and prevention of influenza infection.

- In addition, a contact investigation will be done with people who may have been exposed to the sick person.
- Contact investigations are one of the ways CDC works with partners in the United States and other countries to protect the health of people exposed to an illness. This process involves finding, interviewing, and, in some cases, testing or treating the people who came into contact with the sick person.
- The purpose of a contact investigation for a traveler with H7N9 flu returning from China is to:
  - provide information to exposed passengers and crew so they can get preventive antiviral treatment if needed, recognize any symptoms of disease, isolate themselves if needed, and receive care.
  - refer passengers or crew with influenza-like illness for medical evaluation, testing, and antiviral treatment, if needed.
  - determine whether spread of H7N9 flu may have occurred on the flight and which passengers were at greatest risk.
- All of this can reassure us that the virus is not spreading further in the United States.

#### **Public Health Concern**

- The concern right now is that this H7N9 virus might either mutate or adapt to allow
  efficient transmission during the infection of mammals, or <u>reassort</u> its gene
  segments with human influenza viruses during the co-infection of a single host,
  which would result in a new virus that might be transmissible from person to person.
- Such events are believed to have preceded the influenza pandemics of 1918, 1957, 1968 and 2009.
- Because H7N9 viruses do not commonly infect humans, there is probably little or no immune protection against them in the human population. (CDC is conducting serologic studies to confirm this.)
- If sustained, ongoing human-to-human transmission of H7N9 viruses begins, many more people will become ill, some severely, and unfortunately more deaths will occur.
- CDC is preparing for various scenarios including sustained human-to-human transmission to ensure that the agency will be in the best position to help protect the public from this virus.
- It is important to note, however, that efficient and sustained human-to-human transmission would be needed to trigger a pandemic. There is no indication that such transmission is happening.
- The investigation is ongoing and CDC will provide more information as it becomes available at <a href="http://www.cdc.gov/flu/avianflu/h7n9-virus.htm">http://www.cdc.gov/flu/avianflu/h7n9-virus.htm</a>.

### What CDC Is Doing

- CDC is following this situation closely and coordinating with domestic and international partners, including China CDC and World Health Organization.
- On June 7, 2013, CDC published updated <u>case definitions</u> and <u>interim guidance for specimen collection</u>, <u>processing and testing</u>, and distributed a Health Update via the Health Alert Network that describes the updated guidance on H7N9 testing and reporting of cases. This information is available on the CDC H7N9 web site. (See <u>What Clinicians and Public Health Professionals Should Do</u> for more information.)
- On June 6, CDC published on the CDC H7N9 web site "<u>Interim Risk Assessment and Biosafety Level Recommendations for Working with Influenza A (H7N9) Viruses</u>."
   (See H7N9 Interim Risk Assessment and Biosafety Level Recommendations for more information.)
- CDC is taking routine preparedness measures, including the following:
  - Developing candidate vaccine viruses that could be used to make vaccine if it becomes necessary. (See section <u>H7N9 Candidate Vaccine Viruses</u> for more information.)
  - Sharing CDC-developed test kits that detect the H7N9 virus with other public health laboratories. Information about these test kits is available at <a href="http://www.cdc.gov/flu/avianflu/h7n9-detecting-diagnostics.htm">http://www.cdc.gov/flu/avianflu/h7n9-detecting-diagnostics.htm</a>.
    - CDC began shipping the kits on April 25, 2013. To date, 168 test kits have been shipped domestically (107) and internationally (61). Test kits had been distributed to all 50 states, the District of Columbia and Puerto Rico as of May 28, 2013.
  - Conducting serologic testing to detect antibodies to H7N9 viruses.
    - Studies to see if existing seasonal influenza vaccines may provide any cross-reactive\* antibodies to H7N9 infection are ongoing.
      - \*Cross reactivity here refers to the ability of antibodies developed against a vaccine virus to recognize and neutralize the H7N9 virus.
    - Recently CDC started serological studies that will determine if there is any pre-existing cross-reactive immunity against H7N9 in the current U.S. population.
      - Preliminary results suggest there are very little to no preexisting cross-reactive antibodies against H7N9 in all age groups tested.
    - CDC has also been looking at already-developed H7 candidate vaccine viruses (H7N1, H7N3, H7N7) to see if any of these elicit cross-reactive antibodies to H7N9 infection.
  - Conducting ongoing analysis to determine susceptibility of available H7N9
    viruses to the licensed influenza antiviral drugs, oseltamivir (commercially
    known as Tamiflu®) and zanamivir (Relenza®), as well as to investigational
    antiviral drugs.

- It is possible for an influenza virus to develop resistance to an antiviral drug or class of antiviral drugs.
- Antiviral resistance (or reduced susceptibility) means that a flu virus
  has changed in such a way that, when tested in the laboratory, the
  antiviral drug does not effectively block the virus. This may mean that
  the drug is less effective in treating or preventing illnesses caused by
  the virus.
- Resistance can occur spontaneously, or emerge during the course of antiviral treatment or exposure to antiviral drugs.
- Through antiviral susceptibility testing, CDC Influenza Laboratory and China CDC have found evidence of reduced susceptibility to some antiviral drugs in a small number of H7N9 viruses.
- The H7N9 viruses with reduced susceptibility all have a genetic mutation known as the "R292K substitution."
- This mutation is found in the neuraminidase (NA) gene of the virus and is a known resistance marker that has been associated with reduced susceptibility to the neuraminidase inhibitor (NAI) class of influenza antiviral drugs. (Note: This particular mutation has been observed before in other influenza viruses of subtype N2 and N9.)
- CDC is continuing to investigate whether the R292K substitution confers antiviral resistance among patients infected with H7N9.
- Providing recommendations for travelers as needed.
- Together with partners at ports of entry, CDC staff members are assessing ill travelers returning from affected areas to determine whether any additional public health action is needed.
- CDC also is gathering more information to make a more thorough public health risk assessment.

#### What Clinicians and Public Health Professionals Should Do

- CDC has posted new H7N9 testing recommendations and updated case definitions. They are available, along with other resources for clinicians and health professionals, at <a href="http://www.cdc.gov/flu/avianflu/h7n9-healthprofessionals.htm">http://www.cdc.gov/flu/avianflu/h7n9-healthprofessionals.htm</a>.
- The new information is summarized in a Health Update issued by CDC via the Health Alert Network (HAN) on June 7, 2013. The Health Update is available at <a href="http://emergency.cdc.gov/HAN/han00347.asp">http://emergency.cdc.gov/HAN/han00347.asp</a>
- The Health Update replaces older guidance published on April 5, 2013 in a HAN Health Advisory.
- The primary changes from previous guidance are the following:
  - 1) A new recommendation to test only patients with an appropriate exposure history and severe respiratory illness requiring hospitalization

(In the previous guidance issued on April 5, CDC recommended that all persons with relevant exposure history and illness compatible with influenza, regardless of severity, be tested.)

- 2) A request that only confirmed and probable cases of human infection with H7N9 be reported to CDC.
- The updated guidance reflects the most current epidemiology of H7N9 cases, which indicates that almost all H7N9 human infections have resulted in severe respiratory illness and H7N9 has been found rarely among those with milder disease.
- Clinicians should continue to consider the possibility of H7N9 infection in persons
  presenting with respiratory illness requiring hospitalization and appropriate travel or
  exposure history.
- Confirmed and probable cases of human infection with H7N9 in the United States should be reported to CDC within 24 hours of initial detection. However, state health departments are encouraged to investigate all potential cases of H7N9 infection (as described in the HAN Health Update), in order to determine case status.
- Please refer to <a href="http://www.cdc.gov/flu/avianflu/h7n9-healthprofessionals.htm">http://www.cdc.gov/flu/avianflu/h7n9-healthprofessionals.htm</a> for complete guidance.

### **H7N9 Interim Risk Assessment and Biosafety Level Recommendations**

- On June 6, 2013, CDC released an interim risk assessment and biosafety level recommendations for working with influenza A H7N9 viruses.
- This document explains the basic biosafety considerations, laboratory facility needs and other requirements necessary for handling wild type influenza A H7N9 viruses, and/or synthetic constructs in a laboratory setting. The guidance does not apply to work done with vaccine seed strains.
- An interim risk assessment was performed that used the available information on the characteristics of the H7N9 virus that included: pathogenicity (the ability to cause disease) in birds and people; the ability to grow to high concentrations of virus; the origin (or source) of the virus in nature; and the availability of therapeutic agents. In addition, other general features of influenza A viruses such as how long they remain infectious in the environment and the likely routes of transmission among humans was considered.
- The document describes prerequisites that staff must meet prior to working in laboratories with H7N9.
  - For example, all staff working on the virus should be enrolled in a medical surveillance program and are required to receive a seasonal flu vaccine (unless an absolute medical contraindication exists). Procedures for ongoing medical evaluation of staff, including collection of serum when needed, should be in place.
  - All staff must wear proper respiratory protection, as outlined by the guidance, as well as other personal protective wear, and should receive proper training for use of such equipment.
  - And all staff must avoid contact with domestic or wild birds in accordance with USDA policies when away from the workplace.

- The document also describes the proper laboratory design and containment equipment for working on the virus.
- Recommendations also are provided for decontamination of laboratory work areas.
- The recommendations are based on biosafety levels established in the *Biosafety in Microbiological and Biomedical Laboratories*, 5th Edition, 2009 (BMBL) with added enhancements to protect the worker.

#### Biosafety level requirements regarding work with H7N9 virus

- The document recommends that all "in-vitro" (i.e., cell culture) work with the H7N9 virus be conducted in biosafety level 3 laboratories (BLS-3).
- Animal work involving the H7N9 virus must be conducted in Animal BSL-3 (ABSL-3) laboratories with enhanced practices.
- A BSL-3 facility with specific enhancements includes primary (i.e., laboratory
  equipment designed to protect the lab personnel) and secondary (i.e., facilities
  features in place to protect the environment) barriers as well as special practices to
  protect laboratory workers and the public from accidental exposure to H7N9 viruses.

# Biosafety level recommendations regarding work with diagnostic specimens from patients or animals suspected of H7N9 virus infection.

- The document recommends that processing of samples and/or non-culture based diagnostic testing on clinical specimens from patients and animals with suspected H7N9 virus infection may be performed in a biosafety level 2 (BSL-2) laboratory with enhancements.
- All growth of H7N9 virus in cell culture or embryonated eggs from clinical specimens for virus isolation must be performed in a BSL-3 laboratory.

#### What the Public Should Do

- At this time, no cases of human infection with avian influenza A (H7N9) viruses have been detected in the United States and the virus does not seem to be spreading from person to person.
- Other than the advice for travelers or ill persons below, CDC is not making any additional or special recommendations for public action specific to H7N9.

#### **Travelers**

- Travelers should continue to visit <u>www.cdc.gov/travel</u> or follow <u>@CDCtravel</u> on Twitter for up-to-date information about CDC's travel recommendations.
- CDC does not recommend restricting travel to China at this time.
- Travelers to China should practice hand hygiene, follow food safety practices, and avoid contact with animals.
  - Travelers should wash their hands often or use hand sanitizer. They should try not to touch their eyes, nose, or mouth, except with very clean hands.

- Travelers should eat meats and poultry products, including eggs, only if they have been cooked thoroughly.
- Travelers should avoid touching animals, alive or dead, and should stay away from farms, poultry markets, or other markets where there are live or dead animals.
- Symptoms of H7N9 flu include fever, cough, and shortness of breath. If travelers get sick after returning from China, they should tell their doctors about their recent travel.

#### **H7N9 Candidate Vaccine Viruses**

- So far, CDC has completed development of one H7N9 candidate vaccine virus (CVV) that has been designated "A/Shanghai/2/2013(H7N9)-PR8-IDCDC-RG32A."
- On May 23, WHO issued a statement that the IDCDC-RG32A CVV has passed all relevant safety testing and two-way hemagglutination inhibition (HI) test. (Read more at <a href="http://www.who.int/influenza/vaccines/virus/candidates reagents/summary a h7n9">http://www.who.int/influenza/vaccines/virus/candidates reagents/summary a h7n9</a>
   cvv 20130523.pdf.)
- This CVV was generated by reverse genetics using synthetic hemagglutinin (HA) and neuraminidase (NA) genes.
- CDC used plasmid-based reverse genetics to recover a reassortant virus with the hemagglutinin (HA) and neuraminidase (NA) genes of A/Shanghai/2/2013(H7N9) and the remaining genes from A/Puerto Rico/8/1934(H1N1) in accordance with WHO guidelines for pandemic candidate vaccine virus development.
- CDC has developed other potential CVVs using plasmid-based reverse genetics and is currently evaluating if they have significant advantages.
- No CVVs have been created using conventional reassortment at this time.
- At this time, no decision has been made to mass-produce H7N9 vaccine in the United States.
- The World Health Organization has recommended the use of an A/Anhui/1/2013-like virus for the development of A(H7N9) vaccines for pandemic preparedness purposes. (Read more at <a href="http://www.who.int/influenza/human animal interface/influenza h7n9/ProvisionalRecommendation H7N9 31May13.pdf">http://www.who.int/influenza/human animal interface/influenza h7n9/ProvisionalRecommendation H7N9 31May13.pdf</a>)
- See section <u>H7N9 Vaccine Development</u> for more information.

### **H7N9 Vaccine Development**

 While there is no evidence of sustained (ongoing) person-to-person spread of the H7N9 virus, the Centers for Disease Control and Prevention (CDC), Food and Drug Administration (FDA), National Institutes of Health (NIH) and the Department of Health and Human Services (HHS) are taking standard pandemic preparedness precautions.

- As of today, a decision to produce H7N9 vaccine for a national vaccination response has not been made.
- However, given the number and severity of human illnesses from H7N9 in China, HHS and its partners are taking routine steps to develop H7N9 candidate vaccine viruses and are planning for H7N9 vaccine clinical trials.
- Influenza vaccine production is complex and can be unpredictable. It has many critical and time-sensitive steps; delay at any point during these steps can result in delays in the availability of influenza vaccine.
- It usually takes about six months to produce large quantities of influenza vaccine.
- The development of a high-yield candidate vaccine virus is the first step in developing a vaccine.
  - A candidate vaccine virus is a flu virus that CDC (or one of the other WHO Collaborating Centers) selects and prepares for use by vaccine manufacturers to make a flu vaccine. Candidate vaccine viruses are typically chosen based on their similarity to flu viruses spreading and causing illness in people as well as their ability to grow easily in chicken eggs, which is the method of manufacturing influenza vaccine traditionally used.
  - Without a high-yield candidate vaccine virus, it can be very difficult to manufacture vaccine to protect against a new influenza virus.
  - CDC, NIH, FDA, BARDA (Biomedical Advanced Research and Development Authority in the Office of the Assistant Secretary for Preparedness and Response) and vaccine manufacturers are collaborating to develop high-yield candidate vaccine viruses that could be used to begin production of an H7N9 vaccine.
- BARDA has existing contracts in place with manufacturers of U.S.-licensed influenza vaccines that use egg-, cell-, and recombinant-based technologies. These contracts can support the development, manufacturing, and clinical evaluation of H7N9 vaccines and adjuvants. These contracts also ensure that the candidate vaccine viruses developed would be shared freely and would support commercial scale manufacturing, if needed.
- Once the candidate vaccine virus has been developed, manufacturers can use it to develop a vaccine against the avian influenza A (H7N9) virus.
- Once a vaccine manufacturer receives a candidate vaccine virus, the manufacturer
  then creates what is known as a "seed strain." The seed strain is adapted to make
  the virus grow better using the manufacturer's technology and production systems.
  Once the seed strain is prepared, the vaccine manufacturer uses it to grow large
  quantities of virus for producing flu vaccine.

- Influenza vaccine manufacturers need 8 to 11 weeks to make small lots of vaccine and test whether the candidate vaccine virus works well in the manufacturing process.
  - FDA and other WHO Essential Regulatory Laboratories will make reagents to test the potency of both cell- and egg-based vaccine lots.
- NIH and vaccine manufacturers will sponsor clinical studies of investigational H7N9 vaccines to evaluate the safety and determine the optimal dosing and whether an adjuvant is needed for an adequate immune response. FDA is working closely with NIH and other stakeholders in the design of these clinical trials.
  - The clinical trials will be conducted among people who volunteer to participate in vaccine studies. Information about safety and how people's immune systems respond to the vaccine will be important for planning vaccine production and a vaccine program, if such a program is needed.
- If it is decided that an H7N9 vaccination program is needed, even if production goes as planned, it would be several months before the first doses of H7N9 influenza vaccines are available.
- Although no decision has been made to initiate a H7N9 vaccination program in the
  United States, CDC recommends that local authorities and preparedness programs
  take time to review and update their pandemic influenza vaccine preparedness plans
  since it could take several months to ready a vaccination program, if one were
  necessary.
- Keep in mind that CDC and HHS continue to gather information to make a more thorough public health risk assessment. This is an evolving situation and there is still much to learn.
- Information will be shared as available at <a href="http://www.cdc.gov/flu/avianflu/h7n9-virus.htm">http://www.cdc.gov/flu/avianflu/h7n9-virus.htm</a>.

#### **Avian Influenza Background**

- The H7N9 viruses recently reported in China are the first known human cases of H7N9 influenza infection.
- To date, there have been no human infections with H7N9 in the United States or any countries other than China.
- Different avian influenza A (H7N9) viruses have been identified in birds in North America. Wild waterfowl and shore birds may carry the virus during migrations and may introduce it to domestic poultry. The North American lineage of H7N9 is different from the Eurasian lineage of H7N9 viruses that are currently circulating in China.
- The threat to humans from the North American lineage of H7N9 influenza viruses is low.
- Avian flu viruses do <u>not</u> normally infect humans. However, sporadic human infections with avian flu do occasionally occur.

- Most commonly, human cases of avian influenza happen in people with direct exposure to infected poultry.
- Infected birds can shed a lot of flu virus, for example, in their droppings or their mucus. If someone touches an infected bird or an environment contaminated with virus and then touches their eyes, nose or mouth, they may be infected with bird flu virus.
- There is some evidence that infection may also occur if the flu virus becomes airborne, such as when an infected bird flaps it wings. If someone were to breathe in airborne virus, it's possible they could get infected.
- While most instances of human infection with animal influenza viruses do not result
  in human-to-human transmission, each case should be fully investigated to be sure
  that such viruses are not spreading among humans and to limit further exposure of
  humans to infected animals, if infected animals are identified.
- Poultry, poultry products (eggs) and pork can be safely consumed provided they are properly cooked and properly handled during food preparation.
- Surveillance for avian influenza viruses in North American birds is under the purview
  of the United States Department of Agriculture (USDA) and the Department of the
  Interior (DOI). Questions regarding avian influenza infections in U.S. birds should be
  referred to these agencies.
- International cases of novel influenza A are reportable to the World Health Organization under the International Health Regulations (IHR 2005).
- In 2007, human infection with a novel influenza A virus became a nationally notifiable condition in the United States. Novel influenza A virus infections include all human infections with influenza A viruses that are different from currently circulating human influenza H1 and H3 viruses. Novel viruses include those that are subtyped as non-human in origin and those that are unsubtypable with standard methods and reagents.
- For more information about avian influenza, visit the CDC website at <a href="http://www.cdc.gov/flu/avianflu/index.htm">http://www.cdc.gov/flu/avianflu/index.htm</a>.

#### What is Known about Spread of Other Avian Influenza Viruses, Summary

- Person-to-person spread of other avian influenza viruses is thought to have occurred in the past, most notably with H5N1 viruses.
- In the majority of these instances, spread occurred after prolonged and close contact between the sick person and someone caring for them (most often a family member).
- See "Background on Human Infections with other Avian Influenza Viruses" at <a href="http://www.cdc.gov/flu/avianflu/h5n1-human-infections.htm">http://www.cdc.gov/flu/avianflu/h5n1-human-infections.htm</a> for more information.

#### **Human-to-Human Spread, Background**

- It's important to remember that human-to-human transmission ranges along a continuum; from occasional, "dead-end" human-to-human transmission, to efficient and sustained human-to-human transmission.
- "Dead end" transmission usually refers to when a virus from an animal host infects a person and then there is some subsequent transmission that eventually burns out.
- For example, when a host infects one person who then subsequently infects someone else that is called "first generation spread." If that second person then infects someone else that is called "second generation spread," and so forth.
- Previously, third generation transmission of H5N1 viruses has been documented in one instance at least (Pakistan). (<u>WHO, Weekly Epidemiological Record. "Human</u> <u>cases of avian influenza A(H5N1) in North-West Frontier Province, Pakistan,</u> October-November 2007.")
- However, efficient and sustained (ongoing) transmission in the community is needed for an influenza pandemic to begin.

## **CDC and China, Background**

- U.S. CDC Influenza Division began working with the China National Influenza Center (CNIC), part of the China CDC, in the late 1980s.
- CDC helped China to establish the Chinese National Influenza Surveillance Network and laboratory capability in order to capture more of the influenza viruses circulating in China.
- Since 2004, CDC and China CDC have participated in a series of cooperative
  agreements that have further improved and sustained China's surveillance network
  and supported genetic, antigenic and drug resistance surveillance (in part to inform
  vaccine recommendations), and also strengthened influenza response capacity at all
  levels.
- In October 2010, CNIC was designated as a World Health Organization Collaborating Center for Reference and Research on Influenza.
- CNIC is one of a handful of WHO Collaborating Centers for Reference and Research on Influenza in the world (U.S. CDC in Atlanta, Georgia also is a WHO Collaborating Center).
- Among other things, as a Collaborating Center CNIC regularly provides information from China's recently enhanced surveillance system to help inform decisions about the composition of the seasonal flu vaccine.
- Collaborating Centers also train researchers in specialized techniques, collect epidemiological information on influenza disease prevalence in China and surrounding countries, and assist in developing pandemic preparedness plans. They also receive, characterize, and preserve representative vaccine viruses sent from laboratories around the world, then share that information with other researchers.
- U.S. CDC has an office with an influenza program in China.

- The office includes 1 U.S. Direct Hire and 3 local employees dedicated to the influenza program.
- U.S. CDC in China has a total of 54 staff members, including one secunded to WHO.
- Apart from influenza team staff, other staff with expertise in laboratory, epidemiology and communications have supported the H7N9 response and assisted the Embassy committee in tracking the outbreak.

## **Links to Additional Information**

#### CDC Resources

- CDC will provide updated information as it becomes available at <a href="http://www.cdc.gov/flu/avianflu/h7n9-virus.htm">http://www.cdc.gov/flu/avianflu/h7n9-virus.htm</a>.
- CDC's June 7, 2013, Health Alert Network (HAN) Health Update is available at http://emergencydev.cdc.gov/HAN/han00347.asp
- Other <u>materials for health professionals</u>, <u>laboratorians and clinicians</u> are also available on the CDC H7N9 web site, including:
  - Interim Guidance for Specimen Collection, Processing, and Testing for Patients Who May Be Infected with Avian Influenza A (H7N9) Virus (Posted June 7). <a href="http://www.cdc.gov/flu/avianflu/h7n9/specimen-collection.htm">http://www.cdc.gov/flu/avianflu/h7n9/specimen-collection.htm</a>
  - Interim Guidance on Case Definitions to be Used for Novel Influenza A (H7N9)
     Case Investigations in the United States (Update posted June 7).
     http://www.cdc.gov/flu/avianflu/h7n9/case-definitions.htm
  - Interim Risk Assessment and Biosafety Level Recommendations for Working with Influenza A (H7N9) Viruses (posted June 6). http://www.cdc.gov/flu/avianflu/h7n9/risk-assessment.htm
  - Interim Guidance on the Use of Antiviral Agents for Treatment of Human Infections with Avian Influenza A (H7N9) http://www.cdc.gov/flu/avianflu/h7n9-antiviral-treatment.htm
  - Interim Guidance for Infection Control Within Healthcare Settings When
     Caring for Patients with Confirmed, Probable, or Cases Under Investigation of
     Avian Influenza A(H7N9) Virus Infection
     http://www.cdc.gov/flu/avianflu/h7n9-infection-control.htm
- A new diagram depicting the origins of the H7N9 virus in China is available on the CDC website. The diagram shows how the H7N9 virus's genes are derived from other influenza viruses found in birds. The diagram is available for download from CDC's H7N9 virus images page <a href="http://www.cdc.gov/flu/avianflu/h7n9-images.htm">http://www.cdc.gov/flu/avianflu/h7n9-images.htm</a> and also via the Public Health Image Library (PHIL): <a href="http://phil.cdc.gov/phil/whatsnew.asp">http://phil.cdc.gov/phil/whatsnew.asp</a> (image ID#15798).

- CDC has posted influenza pandemic response planning tips for H7N9 for state and local officials. These tips are available at <a href="http://www.cdc.gov/flu/avianflu/h7n9-additional-information.htm">http://www.cdc.gov/flu/avianflu/h7n9-additional-information.htm</a>.
- Additional resources are available on the CDC H7N9 web site at http://www.cdc.gov/flu/avianflu/h7n9-virus.htm.

#### WHO Resources

- WHO updates information related to human cases of H7N9 avian influenza (and other novel viruses) at <a href="http://www.who.int/csr/don/en/">http://www.who.int/csr/don/en/</a>.
- WHO has posted a Q&A for human infections with H7N9 avian influenza virus. It is available at http://www.who.int/influenza/human animal interface/fag H7N9/en/index.html.
- Information about candidate vaccine viruses is available on the WHO web site at http://www.who.int/influenza/vaccines/virus/en/.
- The World Health Organization Representative Office in China has published the statement and transcript from a press briefing held after the international H7N9 assessment team completed its mission to China. These materials who are available at http://www.wpro.who.int/china/en/.

#### Resources from Other Organizations

- The Chinese Center for Disease Control and Prevention has posted a Q&A document related to this situation. It is available at <a href="http://www.chinacdc.cn/en/research">http://www.chinacdc.cn/en/research</a> 5311/FAQ/201304/t20130418 80053.html.
- The European Centre for Disease Prevention and Control is publishing their latest updates and risk assessments on influenza A(H7N9) at "Avian influenza in humans." This page is available at <a href="http://ecdc.europa.eu/en/healthtopics/avian influenza/whats\_new/Pages/whats\_new/Pages/whats\_new/aspx">http://ecdc.europa.eu/en/healthtopics/avian influenza/whats\_new/Pages/whats\_new/Pages/whats\_new/aspx</a>.